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\*\*\* YOU HAVE NEW MAIL \*\*\*

=> s l1 and conformation

L2 12665 L1 AND CONFORMATION

=> s 12 and reporter

L3 7251 L2 AND REPORTER

=> s 13 and quencher

L4 1381 L3 AND QUENCHER

=> s 14 and monitoring (2a)fluorescence

L5 52 L4 AND MONITORING (2A) FLUORESCENCE

=> s 15 and ratio (5a) intensities

L6 3 L5 AND RATIO (5A) INTENSITIES

=> d 16 bib abs 1-3

L6 ANSWER 1 OF 3 USPATFULL on STN

AN 2003:207233 USPATFULL

TI Nucleic acid **probes** and methods to detect and/or quantify nucleic acid analytes

IN Davies, Martin, Kent, UNITED KINGDOM
Bruce, Ian, East Sussex, UNITED KINGDOM

Wolter, Andreas, Hamburg, GERMANY, FEDERAL REPUBLIC OF

PA PROLIGO, LLC, Boulder, CO, UNITED STATES, 80301 (non-U.S. corporation)

PI US 2003143591 A1 20030731

AI US 2002-278047 A1 20021021 (10)

PRAI US 2001-336432P 20011019 (60)

DT Utility

FS APPLICATION

LREP SWANSON & BRATSCHUN L.L.C., 1745 SHEA CENTER DRIVE, SUITE 330, HIGHLANDS RANCH, CO, 80129

CLMN

Number of Claims: 60

Exemplary Claim: 1 ECLDRWN 21 Drawing Page(s) LN.CNT 3575 CAS INDEXING IS AVAILABLE FOR THIS PATENT. The invention comprises novel methods and strategies to detect and/or quantify nucleic acid analytes. The methods involve nucleic acid probes with covalently conjugated dyes, which are attached either at adjacent nucleotides or at the same nucleotide of the probe and novel linker molecules to attach the dyes to the probes. The nucleic acid probes generate a fluorescent signal upon hybridization to complementary nucleic acids based on the interaction of one of the attached dyes, which is either an intercalator or a DNA groove binder, with the formed double stranded DNA. The methods can be applied to a variety of applications including homogeneous assays, real-time PCR monitoring, transcription assays, expression analysis on nucleic acid microarrays and other microarray applications such as genotyping (SNP analysis). The methods further include pH-sensitive nucleic acid probes that provide switchable fluorescence signals that are triggered by a change in the pH of the medium. CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 2 OF 3 USPATFULL on STN L6 1999:27394 USPATFULL ANHybridization assay using self-quenching fluorescence probe TILivak, Kenneth J., San Jose, CA, United States TN Flood, Susan J. A., Fremont, CA, United States Marmaro, Jeffrey, Aurora, CO, United States Mullah, Khairuzzaman Bashar, Union, CA, United States Perkin-Elmer Corporation, Foster, CA, United States (U.S. corporation) PΑ 19990302 PΤ US 5876930 US 1995-558303 19951115 (8) ΑI Continuation of Ser. No. US 1994-340558, filed on 16 Nov 1994, now RLI patented, Pat. No. US 5538848 DТ Utility Granted FS Primary Examiner: Jones, W. Gary; Assistant Examiner: Riley, Jezia EXNAM Wilson Sonsini Goodrich & Rosati LREP Number of Claims: 39 CLMN ECL Exemplary Claim: 1 5 Drawing Figure(s); 5 Drawing Page(s) DRWN LN.CNT 1413 CAS INDEXING IS AVAILABLE FOR THIS PATENT. A hybridization assay is provided which uses an AB oligonucleotide probe which includes a fluorescent reporter molecule and a quencher molecule capable of quenching the fluorescence of the reporter molecule. The oligonucleotide probe is constructed such that the probe exists in at least one single-stranded conformation when unhybridized where the quencher molecule is near enough to the reporter molecule to quench the fluorescence of the reporter molecule. The oligonucleotide probe also exists in at least one conformation when hybridized to a target polynucleotide where the quencher molecule is not positioned close enough to the reporter molecule to quench the fluorescence of the reporter molecule. By adopting these hybridized and unhybridized

conformations, the reporter molecule and quencher

molecule on the probe exhibits different fluorescence signal intensities

when the probe is hybridized and unhybridized. As a result, it is possible to determine whether the probe is hybridized or unhybridized

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L<sub>6</sub>

based on a change in the fluorescence intensity of the **reporter** molecule, the **quencher** molecule, or a combination thereof. In addition, because the probe can be designed such that the **quencher** molecule quenches the **reporter** molecule when the probe is not hybridized, the probe can be designed such that the **reporter** molecule exhibits limited fluorescence until the probe is either hybridized or digested.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 3 OF 3 USPATFULL on STN

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96:65444 USPATFULL
AN
       Method for detecting nucleic acid amplification using self-quenching
TТ
       fluorescence probe
       Livak, Kenneth J., San Jose, CA, United States
IN
       Flood, Susan J. A., Fremont, CA, United States
       Marmaro, Jeffrey, Foster City, CA, United States
       Applied Biosystems Division, Perkin-Elmer Corp., Foster City, CA, United
PA
       States (U.S. corporation)
PI
       US 5538848
                               19960723
       US 1994-340558
                               19941116 (8)
ΑI
       Utility
DT
       Granted
FS
      Primary Examiner: Jones, W. Gary; Assistant Examiner: Rees, Dianne
EXNAM
       Haynes & Davis
LREP
       Number of Claims: 24
CLMN
       Exemplary Claim: 1
ECL
       1 Drawing Figure(s); 1 Drawing Page(s)
DRWN
LN.CNT 685
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       A method is provided for monitoring the progress of nucleic acid
ΔR
       amplifications that rely on a nucleic acid polymerase having
       5'→3' exonuclease activity. An important feature of the method is
       providing an oligonucleotide probe having a reporter molecule
       and a quencher molecule at either end such that the
       quencher molecule substantially quenches any fluorescence from
       the reporter whenever the oligonucleotide probe is in a single
       stranded state and such that the reporter is substantially
       unquenched whenever the oligonucleotide probe is in a double stranded
       state hybridized to a target polynucleotide.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
=> d his
     (FILE 'HOME' ENTERED AT 16:58:20 ON 13 APR 2004)
     FILE 'BIOSIS, MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 16:58:40 ON
     13 APR 2004
         121246 S PROBES AND HYBRIDIZATION
L1
          12665 S L1 AND CONFORMATION
L2
L3
           7251 S L2 AND REPORTER
L4
           1381 S L3 AND OUENCHER
             52 S L4 AND MONITORING (2A) FLUORESCENCE
L_5
              3 S L5 AND RATIO (5A) INTENSITIES
=> s 15 and intensities (10a) ratio
             5 L5 AND INTENSITIES (10A) RATIO
=> s 17 not 16
             2 L7 NOT L6
```

=> d 18 bib abs 1-2

## L8ANSWER 1 OF 2 USPATFULL on STN AN2001:112052 USPATFULL Detection of nucleic acids by strand displacement TI Nadeau, James G., Chapel Hill, NC, United States IN Hsieh, Helen V., Durham, NC, United States Pitner, J. Bruce, Durham, NC, United States Linn, C. Preston, Durham, NC, United States Becton, Dickinson and Company, Franklin Lakes, NJ, United States (U.S. PΑ corporation) US 6261784 20010717 PΙ US 2000-599164 20000622 (9) AΙ Continuation of Ser. No. US 1999-235583, filed on 22 Jan 1999, now RLI patented, Pat. No. US 6130047 Continuation of Ser. No. US 1997-933749, filed on 23 Sep 1997, now patented, Pat. No. US 5935791 DTUtility GRANTED FS Primary Examiner: Horlick, Kenneth R. EXNAM Highet, David W. LREP Number of Claims: 12 CLMN ECL Exemplary Claim: 1 4 Drawing Figure(s); 4 Drawing Page(s) DRWN CAS INDEXING IS AVAILABLE FOR THIS PATENT. AΒ Detector nucleic acids are employed for detection of nucleic acid target sequences by fluorescence quenching mechanisms. The detector nucleic acid comprises at least two oligonucleotides and is partially single-stranded and partially double-stranded. One of the two dyes of a donor/acceptor dye pair is linked to the first oligonucleotide and the other is linked to a second oligonucleotide such that they are in close spatial proximity when the first and second oligonucleotides are base-paired and donor fluorescence is quenched. A single second oligonucleotide may be hybridized to the first oligonucleotide or multiple second oligonucleotides may be hybridized to the first oligonucleotide and to each other, forming a junction structure comprising multiple donor/acceptor dye pairs. The detector oligonucleotide retains its partially single-stranded and partially double-stranded conformation in the absence of target. In the presence of target, however, the second oligonucleotide(s) of the detector nucleic acid is/are completely or partially displaced from the first, increasing the distance between the donor and acceptor dyes and causing a change in fluorescence which may be detected as an indication of the presence of the target sequence. CAS INDEXING IS AVAILABLE FOR THIS PATENT. rsANSWER 2 OF 2 USPATFULL on STN AN2000:134714 USPATFULL TTDetection of nucleic acids by fluorescence quenching TN Nadeau, James G., Chapel Hill, NC, United States Hsieh, Helen V., Durham, NC, United States Pitner, J. Bruce, Durham, NC, United States Linn, C. Preston, Durham, NC, United States PABeckon, Dickson and Company, Franklin Lakes, NJ, United States (U.S. corporation) US 6130047 PΙ 20001010 ΑТ US 1999-235583 19990122 (9) Continuation of Ser. No. US 1997-933749, filed on 23 Sep 1997, now RLT patented, Pat. No. US 5935791 DTUtility

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FS Granted

EXNAM Primary Examiner: Horlick, Kenneth R.

LREP Fugit, Donna R.

CLMN Number of Claims: 29 ECL Exemplary Claim: 1

DRWN 4 Drawing Figure(s); 4 Drawing Page(s)

LN.CNT 1265

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Detector nucleic acids are employed for detection of nucleic acid target sequences by fluorescence quenching mechanisms. The detector nucleic acid comprises at least two oligonucleotides and is partially single-stranded and partially double-stranded. One of the two dyes of a donor/acceptor dye pair is linked to the first oligonucleotide and the other is linked to a second oligonucleotide such that they are in close spatial proximity when the first and second oligonucleotides are base-paired and donor fluorescence is quenched. A single second oligonucleotide may be hybridized to the first oligonucleotide or multiple second oligonucleotides may be hybridized to the first oligonucleotide and to each other, forming a junction structure comprising multiple donor/acceptor dye pairs. The detector oligonucleotide retains its partially single-stranded and partially double-stranded conformation in the absence of target. In the presence of target, however, the second oligonucleotide(s) of the detector nucleic acid is/are completely or partially displaced from the first, increasing the distance between the donor and acceptor dyes and causing a change in fluorescence which may be detected as an indication of the presence of the target sequence.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.